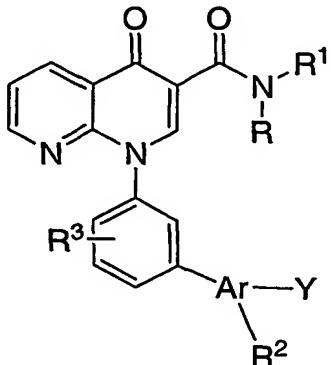


WHAT IS CLAIMED IS:

1. A compound represented by Formula (I):



5

(I)

or a pharmaceutically acceptable salt thereof, wherein

Ar is phenyl, pyridyl, pyrimidyl, indolyl, quinolinyl, thienyl, pyridonyl, oxazolyl, oxadiazolyl, thiadiazolyl, or imidazolyl; or oxides thereof when Ar is a heteroaryl;
 10 Y is -COOR⁴, -C₁₋₆alkyl(C₁₋₄alkyl)_n-COOR⁴, -C₃₋₄cycloalkyl(C₁₋₄alkyl)m-COOR⁴, wherein the -C₁₋₆alkyl and the C₃₋₄cycloalkyl is optionally substituted with halogen, alkoxy, hydroxy or nitrile, and the (C₁₋₄alkyl) substituents are optionally linked to form a C₃₋₄cycloalkyl ; wherein n is 0, 1, 2, 3 or 4, m is 0, 1 or 2;

R and R⁴ are each independently selected from H and -C₁₋₆alkyl;

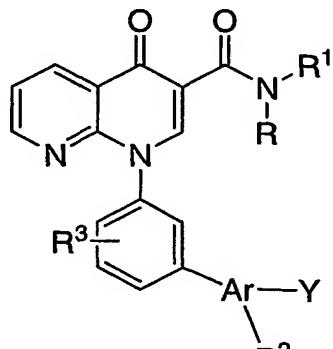
15 R¹ is H, or -C₁₋₆alkyl, -C₃₋₆cycloalkyl, -C₁₋₆alkoxy, -C₂₋₆alkenyl, -C₃₋₆alkynyl, heteroaryl, or heterocycle group, optionally substituted with 1-3 independent haloC₁₋₆alkyl, -C₁₋₆alkyl, -C₁₋₆alkoxy, OH, amino, -(C₀₋₆alkyl)-SO_p-(C₁₋₆alkyl), nitro, CN, =N-O-C₁₋₆alkyl, -O-N=C₁₋₆alkyl, or halogen substituents, wherein p is 0, 1 or 2, or R¹ is C₃₋₆cycloalkyl substituted with phenyl;

20 R² is H, halogen, -CN, -NO₂, -C₁₋₆alkyl, -C₃₋₆cycloalkyl, -O- C₃₋₆cycloalkyl, O-C₁₋₆alkyl, O-C₃₋₆cycloalkyl-C₁₋₆alkyl(C₃₋₆cycloalkyl)(C₃₋₆cycloalkyl), -C₁₋₆alkoxy, phenyl, heteroaryl, heterocycle, amino, -C(O)-C₁₋₆alkyl, -C(O)-O-C₁₋₆alkyl, -C₁₋₆alkyl(=N-OH), -C(N=NOH)C₁₋₆alkyl, -C₀₋₆alkyl(oxy)C₁₋₆alkyl-phenyl, -SO_kNH(C₀₋₆alkyl), or -(C₀₋₆alkyl)-SO_k-(C₁₋₆alkyl), wherein the phenyl, heteroaryl or heterocycle is 25 optionally substituted with halogen, -C₁₋₆alkyl, -C₁₋₆alkoxy, hydroxy, amino, or

-C(O)-O-C₁₋₆alkyl, and wherein the alkyl or cycloalkyl is optionally substituted with 1-6 independently selected halogens or -OH, and wherein k is 0, 1, or 2;

R³ is selected from H, halogen, CN, -C₁₋₆alkyl, -C₃₋₆cycloalkyl, nitro, -C(O)-C₁₋₆alkyl, -C(O)-O-C₀₋₆alkyl, -SO_{n'}NH(C₀₋₆alkyl), or -(C₀₋₆alkyl)-SO_{n'}-(C₁₋₆alkyl), O-C₁₋₆alkyl, O-C₃₋₆cycloalkyl, wherein n' is 0, 1, or 2 and wherein the alkyl and cycloalkyl is optionally substituted with 1-6 independently selected halogen or OH.

2. A compound represented by Formula (I):



10

(I)

or a pharmaceutically acceptable salt thereof, wherein

Ar is phenyl, pyridyl, pyrimidyl, indolyl, quinolinyl, thienyl, pyridonyl,
15 oxazolyl, oxadiazolyl, thiadiazolyl, or imidazolyl; or oxides thereof when Ar is a heteroaryl;
Y is -COOH, -C₁₋₆alkyl(C₁₋₄alkyl)_n-COOH, -C₃₋₄cycloalkyl(C₁₋₄alkyl)_m-COOH, wherein the -C₁₋₆alkyl and the C₃₋₄cycloalkyl is optionally substituted with halogen, alkoxy, hydroxy or nitrile, and the (C₁₋₄alkyl) substituents are optionally linked to form a C₃₋₄cycloalkyl ; wherein n is 0, 1, 2, 3 or 4, m is 0, 1 or 2;

20 R is H or -C₁₋₆alkyl;

R¹ is H, or -C₁₋₆alkyl, -C₃₋₆cycloalkyl, -C₁₋₆alkoxy, -C₂₋₆alkenyl, -C₃₋₆alkynyl, heteroaryl, or heterocycle group, optionally substituted with 1-3 independent haloC₁₋₆alkyl, -C₁₋₆alkyl, -C₁₋₆alkoxy, OH, amino, -(C₀₋₆alkyl)-SO_p-(C₁₋₆alkyl), nitro, CN, =N-O-C₁₋₆alkyl, -O-N=C₁₋₆alkyl, or halogen substituents, wherein p is 0, 1 or 2;

R² is H, halogen, -CN, -NO₂, -C₁₋₆alkyl, -C₃₋₆cycloalkyl, -O- C₃₋₆cycloalkyl, O-C₁₋₆alkyl, O-C₃₋₆cycloalkyl-C₁₋₆alkyl(C₃₋₆cycloalkyl)(C₃₋₆cycloalkyl), -C₁₋₆alkoxy, phenyl, heteroaryl, heterocycle, amino, -C(O)-C₁₋₆alkyl, -C(O)-O-C₁₋₆alkyl, -C₁₋₆alkyl(=N-OH), -C(N=NOH)C₁₋₆alkyl, -C₀₋₆alkyl(oxy)C₁₋₆alkyl-phenyl, -SO_kNH(C₀₋₆alkyl), or -(C₀₋₆alkyl)-SO_k-(C₁₋₆alkyl), wherein the phenyl, heteroaryl or heterocycle is

5 optionally substituted with halogen, -C₁₋₆alkyl, -C₁₋₆alkoxy, hydroxy, amino, or -C(O)-O-C₁₋₆alkyl, and wherein the alkyl or cycloalkyl is optionally substituted with 1-6 independently selected halogens or -OH, and wherein k is 0, 1, or 2;

R³ is selected from H, halogen, CN, -C₁₋₆alkyl, -C₃₋₆cycloalkyl, nitro, -

10 C(O)-C₁₋₆alkyl, -C(O)-O-C₀₋₆alkyl, -SO_{n'}NH(C₀₋₆alkyl), or -(C₀₋₆alkyl)-SO_{n'}-(C₁₋₆alkyl), O-C₁₋₆alkyl, O-C₃₋₆cycloalkyl, wherein n' is 0, 1, or 2 and wherein the alkyl and cycloalkyl is optionally substituted with 1-6 independently selected halogen or OH.

15 3. The compound according to claim 2, or a pharmaceutically acceptable salt thereof, wherein

Y is -C₃₋₄cycloalkyl(C₁₋₄alkyl)_m-COOH, wherein the C₃₋₄cycloalkyl is optionally substituted with halogen, alkoxy, hydroxy or nitrile, and the (C₁₋₄alkyl) substituents are optionally linked to form a C₃₋₄cycloalkyl ; wherein n is 0, 1, 2, 3 or 4, m is 0, 1 or 2.

20

4. The compound according to claim 2, or a pharmaceutically acceptable salt thereof, wherein

Y is cyclopropyl-COOH;

Ar is phenyl.

25

5. The compound according to claim 4, or a pharmaceutically acceptable salt thereof, wherein

30 R¹ is -C₁₋₆alkyl optionally substituted with 1-3 independent -C₁₋₆alkyl, -C₁₋₆alkoxy, OH, amino, -(C₀₋₆alkyl)-SO_p-(C₁₋₆alkyl), nitro, CN, =N-O-C₁₋₆alkyl, -O-N=C₁₋₆alkyl, or halogen substituents.

6. The compound according to claim 4, or a pharmaceutically acceptable salt thereof, wherein

R^1 is $-C_3\text{-}6\text{cycloalkyl}$ optionally substituted with 1-3 independent $-C_1\text{-}6\text{alkyl}$, $-C_1\text{-}6\text{alkoxy}$, OH, amino, $-(C_0\text{-}6\text{alkyl})-\text{SO}_p-(C_1\text{-}6\text{alkyl})$, nitro, CN, $=N\text{-}O\text{-}C_1\text{-}6\text{alkyl}$, $-O\text{-}N=C_1\text{-}6\text{alkyl}$, or halogen substituents.

5 7. The compound according to claim 4, or a pharmaceutically acceptable salt thereof, wherein

R is hydrogen.

10 8. The compound according to claim 4, or a pharmaceutically acceptable salt thereof, wherein

R^2 is hydrogen or $-C_1\text{-}3\text{alkyl}$.

15 9. The compound according to claim 4, or a pharmaceutically acceptable salt thereof, wherein

R^1 is $-C_3\text{-}6\text{cycloalkyl}$ optionally substituted with methyl or halo; and

R is hydrogen.

20 10. The compound according to claim 4, or a pharmaceutically acceptable salt thereof, wherein

R^1 is cyclopropyl optionally substituted with methyl or halo; and

R and R^2 are hydrogen.

25 11. The compound according to claim 2, or a pharmaceutically acceptable salt, wherein

Ar is pyridyl, pyrimidyl, or oxide thereof.

30 12. The compound according to claim 11, or a pharmaceutically acceptable salt, wherein

R^1 is $-C_1\text{-}6\text{alkyl}$ optionally substituted with 1-3 independent $-C_1\text{-}6\text{alkyl}$, $-C_1\text{-}6\text{alkoxy}$,

OH , amino, $-(C_0\text{-}6\text{alkyl})-\text{SO}_p-(C_1\text{-}6\text{alkyl})$, nitro, CN, $=N\text{-}O\text{-}C_1\text{-}6\text{alkyl}$, $-O\text{-}N=C_1\text{-}6\text{alkyl}$, or halogen substituents.

13. The compound according to claim 11, or a pharmaceutically acceptable salt thereof, wherein

R¹ is -C₃₋₆cycloalkyl optionally substituted with 1-3 independent -C₁₋₆alkyl, -C₁₋₆alkoxy, OH, amino, -(C₀₋₆alkyl)-SO_p-(C₁₋₆alkyl), nitro, CN, =N-O-C₁₋₆alkyl, -O-N=C₁₋₆alkyl, or halogen substituents.

14. The compound according to claim 11, or a pharmaceutically acceptable salt thereof, wherein

R is hydrogen.

10

15. The compound according to claim 11, or a pharmaceutically acceptable salt thereof, wherein

R² is hydrogen or -C₁₋₃alkyl or halogen.

15

16. The compound according to claim 11, or a pharmaceutically acceptable salt thereof, wherein

R¹ is -C₃₋₆cycloalkyl optionally substituted with methyl or halo; and

R is hydrogen.

20

17. The compound according to claim 11, or a pharmaceutically acceptable salt thereof, wherein

R¹ is cyclopropyl optionally substituted with methyl or halo; and

R and R² are hydrogen or halogen;

R³ is hydrogen or halogen.

25

18. The compound according to claim 2, or a pharmaceutically acceptable salt thereof, wherein

R and R³ are hydrogen,;

R¹ is -C₃₋₆cycloalkyl optionally substituted with methyl or halo, or

30 -C₁₋₃alkyl optionally substituted with 1-3 halo; and

Ar is phenyl.

19. The compound according to claim 18 wherein

R² is hydrogen or halo; and

Y is -CH₃-C₃₋₄cycloalkyl-COOH or -C₃₋₄cycloalkyl-COOH.

20. The compound according to claim 2, which is

5 2-(trans)-{3'-[3-[cyclopropylamino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

2-(trans)-{3'-[3-[cyclopropylamino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-3-yl}cyclopropanecarboxylic acid;

10 2-{3'-[3-[cyclopropylamino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-3-yl}-2-methylpropanoic acid;

2-{3'-[3-[cyclopropylamino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}-2-methylpropanoic acid;

15 3-{3'-[3-[cyclopropylamino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}-3-methylbutanoic acid;

1-{3'-[3-[cyclopropylamino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}(hydroxy)acetic acid;

20 1-{3'-[3-[cyclopropylamino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

5-{3'-[3-[cyclopropylamino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}-2,2-dimethyl-1,3-dioxolane-4-carboxylic acid;

1-{3'-[3-[cyclopropylamino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-3-yl}cyclopropanecarboxylic acid;

25 1-cyano-3-{3'-[3-[cyclopropylamino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}-2,2-dimethylcyclopropanecarboxylic acid;

2-(trans)-{3'-[3-[cyclopropylamino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-3-fluoro-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

(cis)-2-{3'-[3-[cyclopropylamino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-30 3-yl}cyclopropanecarboxylic acid;

2-(trans)-{3'-bromo-5'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

2-(trans)-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]-3-methyl-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

5 2-(trans)-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]-2-methyl-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

2-(trans)-{3-chloro-3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

2-(cis)-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]-3-fluoro-1,1'-10 biphenyl-4-yl}cyclopropanecarboxylic acid;

3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]-1,1'-biphenyl-4-carboxylic acid;

2-(trans)-{3'-[3-(morpholin-4-ylcarbonyl)-4-oxo-1,8-naphthyridin-1(4*H*)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

15 2-(trans)-{3'-[4-oxo-3-({[5-(trifluoromethyl)-1,3,4-thiadiazol-2-yl]amino}carbonyl)-1,8-naphthyridin-1(4*H*)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

2-(trans)-{3'-[3-({[2-(methylthio)ethyl]amino}carbonyl)-4-oxo-1,8-naphthyridin-1(4*H*)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

2-(trans)-{3'-[3-({[2-(methylsulfonyl)ethyl]amino}carbonyl)-4-oxo-1,8-naphthyridin-1(4*H*-20 -yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

2-(trans)-{3'-[4-oxo-3-[(2,2,2-trifluoroethyl)amino]carbonyl]-1,8-naphthyridin-1(4*H*)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

2-(trans)-(5-{3-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]phenyl}thien-2-yl)cyclopropanecarboxylic acid;

25 2-(trans)-{3'-[3-[(cyclopropylmethyl)amino]carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid;

2-(trans)-{3'-[3-[(1-cyanocyclopropyl)amino]carbonyl]-4-oxo-1,8-naphthyridin-1(4*H*)-yl]-1,1'-biphenyl-4-yl}cyclopropanecarboxylic acid; or

3-{3'-[3-[(isopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]-1,1'-biphenyl-4-yl}-3-methylbutanoic acid.

21. A compound of claim 1 which is

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(+)-(trans)-2-{3-fluoro-3'-[4-oxo-3-[(2,2,2-trifluoroethyl)amino]carbonyl]-1,8-naphthyridin-1(4H)-yl)biphenyl-4-yl}cyclopropanecarboxylic acid;

1-(3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl)methyl)cyclobutanecarboxylic acid;

10 (trans)-2-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}-2-methylcyclopropanecarboxylic acid;

(trans)-2-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-2-yl}cyclopropanecarboxylic acid;

15 3-methyl-3-{3'-[4-oxo-3-[(2,2,2-trifluoroethyl)amino]carbonyl]-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}butanoic acid;

(trans)-2-{3'-[4-oxo-3-[(2,2,2-trifluoroethyl)amino]carbonyl]-1,8-naphthyridin-1(4H)-yl]biphenyl-2-yl}cyclopropanecarboxylic acid;

(trans)-2-{3'-[4-oxo-3-[(2,2,3,3,3-pentafluoropropyl)amino]carbonyl]-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

20 (trans)-2-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}-1-fluorocyclopropanecarboxylic acid;

(+)-(trans)-2-{3-chloro-3'-(3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

25 (-)-(trans)-2-{3'-[4-oxo-3-[(2,2,2-trifluoroethyl)amino]carbonyl]-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

(+)-(trans)-ethyl 2-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylate;

(+)-(trans)-isopropyl 2-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylate;

tert-butyl 3-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}-2,2-dimethylpropanoate;

3-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}-2,2-dimethylpropanoic acid;

5 3-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-3-yl}-2,2-dimethylpropanoic acid;

1-(3'-(3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl)biphenyl-3-yl)methyl)cyclobutanecarboxylic acid;

3-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-2-yl}-2,2-

10 dimethylpropanoic acid;

1-(3'-(3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl)biphenyl-2-yl)methyl)cyclobutanecarboxylic acid;

(+)-(trans)-2-{3'-(3-[(*tert*-butylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl)biphenyl-4-yl}cyclopropanecarboxylic acid;

15 (+)-(trans)-2-{3'-(3-[(cyclobutylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl)biphenyl-4-yl}cyclopropanecarboxylic acid;

3-{3'-(3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl)biphenyl-4-yl}bicyclo[1.1.1]pentane-1-carboxylic acid;

4-{3'-(3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl)biphenyl-4-yl}-4-

20 hydroxypentanoic acid;

(trans)-2-{3'-(3-{[(\pm)-cis-(2-fluorocyclopropyl)amino]carbonyl}-4-oxo-1,8-naphthyridin-1(4H)-yl)-(+)-biphenyl-4-yl}cyclopropanecarboxylic acid;

(+)-(trans)-2-{3'-(3-{[(dicyclopropylmethyl)amino]carbonyl}-4-oxo-1,8-naphthyridin-1(4H)-yl)biphenyl-4-yl}cyclopropanecarboxylic acid;

25 4-{3'-(3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl)biphenyl-4-yl}-2,2-dimethylbutanoic acid;

(+)-(trans)-2-{3'-(3-{(1-hydroxycyclopropyl)amino]carbonyl}-4-oxo-1,8-naphthyridin-1(4H)-yl)biphenyl-4-yl}cyclopropanecarboxylic acid;

(+)-(trans)-2-{3'-[4-oxo-3-[(1-phenylcyclopropyl)amino]carbonyl]-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

4-{3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}-3,3-dimethylbutanoic acid;

5 (+)-(trans)-2-{3'-[3-[(1-cyclopropyl-1-methylethyl)amino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

1-(3'-[4-oxo-3-[(2,2,2-trifluoroethyl)amino]carbonyl]-1,8-naphthyridin-1(4H)-yl)biphenyl-4-yl)methyl)cyclobutanecarboxylic acid;

(+)-(trans)-2-{3'-[3-[(cyclopropylmethyl)amino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

10 (-)-(trans)-2-{3-fluoro-3'-[3-[(1-hydroxycyclopropyl)amino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

(trans)-2-{3'-[4-oxo-3-[(\pm)-2,2,2-trifluoro-1-methylethyl)amino]carbonyl]-1,8-naphthyridin-1(4H)-yl}-(+)-biphenyl-4-yl}cyclopropanecarboxylic acid;

15 (+)-(trans)-2-{3'-[3-[(1-methylcyclopropyl)amino]carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid;

2,2-dimethyl-4-{3'-[4-oxo-3-[(2,2,2-trifluoroethyl)amino]carbonyl]-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}butanoic acid;

2,2-dimethyl-3-{3'-[4-oxo-3-[(2,2,2-trifluoroethyl)amino]carbonyl]-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}propanoic acid;

20 (-)-(trans)-2-{3-chloro-3'-[3-[(cyclopropylamino)carbonyl]-4-oxo-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid; or

(+)-(trans)-2-{3'-[4-oxo-3-[(2,2,2-trifluoroethyl)amino]carbonyl]-1,8-naphthyridin-1(4H)-yl]biphenyl-4-yl}cyclopropanecarboxylic acid.

25

22. A pharmaceutical composition comprising
a therapeutically effective amount of the compound according to
any one of claims 1 to 21, or a pharmaceutically acceptable salt thereof; and
a pharmaceutically acceptable carrier.

23. The pharmaceutical composition according to claim 22, further comprising a Leukotriene receptor antagonist, a Leukotriene biosynthesis inhibitor, an M2/M3 antagonist, a corticosteroid, an H1 receptor antagonist or a beta 2 adrenoceptor agonist.

5

24. The pharmaceutical composition according to claim 22, further comprising a COX-2 selective inhibitor, a statin, or an NSAID.

25. A method of treatment or prevention of asthma, chronic bronchitis,
10 chronic obstructive pulmonary disease (COPD), eosinophilic granuloma, psoriasis and other benign or malignant proliferative skin diseases, endotoxic shock (and associated conditions such as laminitis and colic in horses), septic shock, ulcerative colitis, Crohn's disease, reperfusion injury of the myocardium and brain, inflammatory arthritis, osteoporosis, chronic glomerulonephritis, atopic dermatitis, urticaria, adult respiratory distress syndrome, infant
15 respiratory distress syndrome, chronic obstructive pulmonary disease in animals, diabetes insipidus, allergic rhinitis, allergic conjunctivitis, vernal conjunctivitis, arterial restenosis, atherosclerosis, neurogenic inflammation, pain, cough, rheumatoid arthritis, ankylosing spondylitis, transplant rejection and graft versus host disease, hypersecretion of gastric acid, bacterial, fungal or viral induced sepsis or septic shock, inflammation and cytokine-mediated
20 chronic tissue degeneration, osteoarthritis, cancer, cachexia, muscle wasting, depression, memory impairment, monopolar depression, acute and chronic neurodegenerative disorders with inflammatory components, Parkinson disease, Alzheimer's disease, spinal cord trauma, head injury, multiple sclerosis, tumour growth and cancerous invasion of normal tissues comprising the step of administering a therapeutically effective amount, or a prophylactically
25 effective amount, of the compound according to claim 1 or a pharmaceutically acceptable salt thereof.

26. A method of enhancing cognition in a healthy subject comprising administering a safe cognition enhancing amount of compound according to claim 1, or a
30 pharmaceutically salt thereof.

27. A method according to claim 25 wherein the amount of compound is insufficient to emisis in said subject.

28. A method according to claim 25 wherein the subject is a human of age 55 or older.

29. A compound according to claim 2 wherein Y is -C₃₋₆ 5 cycloalkyl(C₁₋₄alkyl)_m-COOH, wherein the C₃₋₆cycloalkyl is optionally substituted with halogen, alkoxy, hydroxyl or nitrile, and the (C₁₋₄alkyl) substituents are optionally linked to form a C₃₋₆cycloalkyl; wherein n is 0, 1, 2, 3 or 4, m is 0, 1.

10 30. Use of a compound of Formula (I) as defined in any one of claims 1 to 21 or 29, in the manufacture of a medicament for treatment or prevention of an ailment as set forth in claim 25 or enhancing cognition in a healthy subject.

15 31. A compound of Formula (I) as defined in any one of claims 1 to 21 or 29, for use in treatment or prevention of an ailment as set forth in claim 25 or enhancing cognition in a healthy subject.

32. A phosphodiesterase-4 inhibitor pharmaceutical 20 composition comprising an acceptable phosphodiesterase-4 inhibiting amount of a compound of formula (I) as defined in any one of claims 1 to 21 or 29, in association with a pharmaceutically acceptable carrier.